RECEIVED **CENTRAL FAX CENTER**

ISIS PATENT DEPT

NOV 0 4 2005 PTO/SB/21 (09-04) Approved for use through 07/31/2006. OMB 0631-0031
U.S. Patent and Trademark Office; U.S. DEPARTMENT OF COMMERCE Under the Paperwork Reduction Act of 1995, no persons are required to reasond to a collection of information unless it displays a valid OMB control number. Application Number 10/038.335 Filing Date TRANSMITTAL 01/02/2002

First Named Inventor Ronnie C. Hanecak FORM Art Unit 1636 Examiner Name David Guzo (to be used for all correspondence after initial filing) Attorney Docket Number ISIS-4976 Total Number of Pages In This Submission **ENCLOSURES** (Check all that apply) After Allowance Communication to TC Fee Transmittal Form Drawing(s) Appeal Communication to Board Licensing-related Papers of Appeals and Interferences Fee Attached Appeal Communication to TC 1 Patition (Appeal Notice, Brief, Reply Brief) Amendment/Reply Pelition to Convert to a Proprietary Information Provisional Application After Final Power of Attorney, Revocation Change of Correspondence Address Status Letter Affidavits/declaration(s) Other Enclosure(s) (please identify Terminal Disclaimer Extension of Time Request below): Request for Refund Express Abandonment Request CD. Number of CD(s) Information Disclosure Statement Landscape Table on CD Certified Copy of Priority Remarks Document(s) Reply to Missing Parts/ Incomplete Application Reply to Missing Parts under 37 CFR 1.52 or 1.53 SIGNATURE OF APPLICANT, ATTORNEY, OR AGENT Firm Name Isis Pharmaceuticals, Inc. Signature Printed name Joshua McLaughlin Reg. No. Date 50,455 11/04/2005 CERTIFICATE OF TRANSMISSION/MAILING I hereby certify that this correspondence is being facsimile transmitted to the USPTO or deposited with the United States Postal Service with sufficient postage as first class mail in an envelope addressed to: Commissioner for Patents. P.O. Box 1450, Alexandria, VA 22313-1450 on the date shown below: Signature Date 11/04/2005 Jamie Todpa Typed or printed name

This collection of information is required by 37 CFR 1.5. The information is required to obtain or retain a benefit by the public which is to fits (and by the USPTO to This collection of information is required by 37 CPR 1.5. The information is required to obtain or retain a benefit by the public which is do his lead by the USP 10 to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.11 and 1.14. This collection is estimated to 2 hours to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the bidMoula case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Palent and Trademark Office. U.S. Department of Commerce, P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

If you need essistance in completing the form, cell 1-800-PTO-9189 and select option 2.

RECEIVED CENTRAL FAX CENTER

NOV 0 4 2005

Approved for use through 07/31/2006, OMB 0651-0032

U.S. Patent and Trademark Officer, U.S. DEPARTMENT OF COMMERCE and to a collection of Information unless it diantages a valid OMB control number.

| | Complete if Known | | |
|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------|-------------------------------------------|-----------------------------------|
| Fees pursuant to the Consolidated Appropriations Act. 2005 (H.R. 4818). FEE TRANSMITTAL For FY 2005 | Application Number 10/038,335 | | |
| | Filing Date | 01/02/2002 | |
| | First Named Inventor | Ronnie C. Hanecak | |
| | Examiner Name | David Guzo | |
| Applicant claims small entity status. See 37 CFR 1.27 | Art Unit | 1636 | |
| TOTAL AMOUNT OF PAYMENT (\$) 65.00 | Attorney Docket No. | ISIS-4976 | |
| | | | |
| METHOD OF PAYMENT (check all that apply) | | | |
| Check Credit Card Money Order None Other (please identify): | | | |
| Deposit Account Deposit Account Number: 50-0252 Deposit Account Name: Isis Pharmaceuticals, Inc. | | | |
| For the above-identified deposit account, the Director is hereby authorized to: (check all that apply) | | | |
| Charge fee(s) Indicated below Charge fee(s) indicated below, except for the filing fee | | | |
| | | | |
| under 37 CFR 1 18 and 1 17 | | | |
| WARNING: Information on this form may become public. Credit card information should not be included on this form. Provide credit card information and authorization on PTO-2038. | | | |
| FEE CALCULATION | | | |
| 1. BASIC FILING, SEARCH, AND EXAMINATION FEES | | | |
| FILING FEES SEAI Small Entity | | MINATION FEES Small Entity | |
| Application Type Fee (\$) Fee (\$) | Small Entity Fee (\$) Fee | | Fees Paid (5) |
| Utility 300 150 500 | 250 20 | 0 100 _ | |
| Design 200 100 100 | 50 130 | D 65 - | |
| Plant 200 100 300 | 150 160 | 0 80 - | |
| Reissue 300 150 500 | 250 60 | 0 300 - | |
| Provisional 200 100 0 | 0 | 0 0 - | |
| 2. EXCESS CLAIM FEES | | | Small Entity Fee (\$) Fee (\$) |
| Fee Description Each claim over 20 or, for Reissues, each claim over 20 and more than in the original patent 50 25 | | | |
| Each independent claim over 3 or, for Reissues, each independent claim more than in the original patent 200 100 | | | |
| Multiple dependent claims | manda servici | -l- Danisadad Claima | 360 180 |
| Total Claims Extra Claims Fee (\$) Fee - 20 or HP = x = | | ple Dependent Claims a (\$) Fee Paid (| (5) |
| HP = highest number of total claims paid for, if greater than 20 | | | |
| Indep. Claims Extra Claims Fea (\$) Fea (\$) | Paid (\$) | | |
| HP = highest number of independent claims paid for, if greater than 3 | | | |
| 3, APPLICATION SIZE FEE | | | |
| If the specification and drawings exceed 100 sheets of paper, the application size fee due is \$250 (\$125 for small entity) for each additional 50 sheets or fraction thereof. See 35 U.S.C. 41(a)(1)(G) and 37 CFR 1.16(s). | | | |
| | th additional 50 or fracti | | Fee Paid (\$) |
| 100 = / 50 = (round up to a whole number) x = | | | |
| 4. OTHER FEE(S) Fees Paid (S) | | | |
| Non-English Specification, \$130 fee (no small entity discount) | | | |
| Other: Terminal disclaimer fee under 37 CFR 1.20(d) 65.00 | | | |
| SUBMITTED BY | ······································ | | |
| Signature | Registration No. 50,455 (Attorney/Agent) | Telephone 7 | 760-603-2767 |
| Name (Print/Type) Joshua McLaughlin | f. manualtudent) | Date · //- | |

This collection of information is required by 37 CFR 1.136. The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.14. This collection is estimated to take 30 minutes to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS, SEND TO: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

If you need assistance in completing the form, call 1-800-PTO-9199 and select option 2.